

File Copy  
09/105117

(FILE 'HOME' ENTERED AT 18:56:31 ON 19 OCT 2001)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, SCISEARCH' ENTERED AT 18:57:06 ON  
19 OCT 2001

L1 2893 S MICROBIAL (W) PRODUCTION  
L2 190 S L1 AND (AMINO (W) ACIDS)  
L3 28 S L2 AND CORYNEBACTERIUM  
L4 13 L3 AND LYSINE  
L5 0 L3 AND ((EXPORT) (W) (GENE OR CARRIER))  
L6 0 S L3 AND EXPORT (W) GENE  
L7 166 S EXPORT (W) GENE  
L8 0 S L3 AND L7  
L9 0 S L3 (P) L7  
L10 0 S L2 AND L7  
L11 0 S L2 AND EXPORT (W) GENE  
L12 61 S L7 AND MICROB?  
L13 1 S L12 AND CORYNEBACTERIUM  
L14 26 DUP REM L3 (2 DUPLICATES REMOVED)  
L15 13 DUP REM L4 (0 DUPLICATES REMOVED)

=> log off y

STN  
Search Strategy

File Copy  
09/105117  
EAST  
Search Strategy

Type	L #	Hits	Search Text	DBs	Time Stamp	Comments	Error Definition	Errors
1 BRS	L1	0	microbial adj production adj of adj amino adj acid	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:38			0
2 BRS	L7	702	microbial adj production	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:38			0
3 BRS	L13	314	L7 and (amino adj acid)	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:39			0
4 BRS	L19	47	L13 and Corynebacterium	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:40			0

	Type	L #	Hits	Search Text	DBs	Time Stamp	Comments	Error Definition	Errors
5	BRS	L25	0	L19 and (manufacturing adj L-lysine)	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:42			0
6	BRS	L37	1	L19 and ((export)adj(gene or carrier))	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:47			0
7	BRS	L31	11	L19 and lysine	USPA T; US-P GPUB ; EPO; JPO; DERW ENT	2001/10/1 9 18:48			0

155

File Copy  
09/105117

Connecting via Winsock to STN

Trying 3106016892...Open

Welcome to STN International! Enter x:x

LOGINID:SSSPTA1600RXM

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

\* \* \* \* \* Welcome to STN International \* \* \* \* \*

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America  
NEWS 2 Dec 17 The CA Lexicon available in the CAPLUS and CA files  
NEWS 3 Feb 06 Engineering Information Encompass files have new names  
NEWS 4 Feb 16 TOXLINE no longer being updated  
NEWS 5 Apr 23 Search Derwent WPINDEX by chemical structure  
NEWS 6 Apr 23 PRE-1967 REFERENCES NOW SEARCHABLE IN CAPLUS AND CA  
NEWS 7 May 07 DGENE Reload  
NEWS 8 Jun 20 Published patent applications (A1) are now in USPATFULL  
NEWS 9 JUL 13 New SDI alert frequency now available in Derwent's  
DWPI and DPCI  
NEWS 10 Aug 23 In-process records and more frequent updates now in  
MEDLINE  
NEWS 11 Aug 23 PAGE IMAGES FOR 1947-1966 RECORDS IN CAPLUS AND CA  
NEWS 12 Aug 23 Adis Newsletters (ADISNEWS) now available on STN  
NEWS 13 Sep 17 IMSworld Pharmaceutical Company Directory name change  
to PHARMASEARCH  
NEWS 14 Oct 09 Korean abstracts now included in Derwent World Patents  
Index  
NEWS 15 Oct 09 Number of Derwent World Patents Index updates increased  
NEWS 16 Oct 15 Calculated properties now in the REGISTRY/ZREGISTRY File  
  
NEWS EXPRESS August 15 CURRENT WINDOWS VERSION IS V6.0c,  
CURRENT MACINTOSH VERSION IS V6.0 (ENG) AND V6.0J (JP),  
AND CURRENT DISCOVER FILE IS DATED 07 AUGUST 2001  
NEWS HOURS STN Operating Hours Plus Help Desk Availability  
NEWS INTER General Internet Information  
NEWS LOGIN Welcome Banner and News Items  
NEWS PHONE Direct Dial and Telecommunication Network Access to STN  
NEWS WWW CAS World Wide Web Site (general information)

Enter NEWS followed by the item number or name to see news on that specific topic.

All use of STN is subject to the provisions of the STN Customer agreement. Please note that this agreement limits use to scientific research. Use for software development or design or implementation of commercial gateways or other similar uses is prohibited and may result in loss of user privileges and other penalties.

\* \* \* \* \* STN Columbus \* \* \* \* \*

FILE 'HOME' ENTERED AT 18:56:31 ON 19 OCT 2001

=> file biosis caplus embase medline scisearch		
COST IN U.S. DOLLARS	SINCE FILE	TOTAL
	ENTRY	SESSION
FULL ESTIMATED COST	0.15	0.15

FILE 'BIOSIS' ENTERED AT 18:57:06 ON 19 OCT 2001  
COPYRIGHT (C) 2001 BIOSIS(R)

FILE 'CAPLUS' ENTERED AT 18:57:06 ON 19 OCT 2001  
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.  
PLEASE SEE "HELP USAGETERMS" FOR DETAILS.  
COPYRIGHT (C) 2001 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'EMBASE' ENTERED AT 18:57:06 ON 19 OCT 2001  
COPYRIGHT (C) 2001 Elsevier Science B.V. All rights reserved.

FILE 'MEDLINE' ENTERED AT 18:57:06 ON 19 OCT 2001

FILE 'SCISEARCH' ENTERED AT 18:57:06 ON 19 OCT 2001  
COPYRIGHT (C) 2001 Institute for Scientific Information (ISI) (R)

=> s microbial (w) production  
L1 2893 MICROBIAL (W) PRODUCTION

=> s L1 and (amino (w) acids)  
L2 190 L1 AND (AMINO (W) ACIDS)

=> s L2 and Corynebacterium  
L3 28 L2 AND CORYNEBACTERIUM

=> L3 and lysine  
L4 13 L3 AND LYSINE

=> L3 and ((export) (w) (gene or carrier))  
L5 0 L3 AND ((EXPORT) (W) (GENE OR CARRIER))

=> s L3 and export (w) gene  
L6 0 L3 AND EXPORT (W) GENE

=> s export (w) gene  
L7 166 EXPORT (W) GENE

=> s L3 and L7  
L8 0 L3 AND L7

=> s L3 (p) L7  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L13 (P) L37'  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L14 (P) L38'  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L15 (P) L39'  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L16 (P) L40'  
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH  
FIELD CODE - 'AND' OPERATOR ASSUMED 'L17 (P) L41'  
L9 0 L3 (P) L7

```
=>
=> s L2 and L7
L10          0 L2 AND L7

=> s L2 and export (w) gene
L11          0 L2 AND EXPORT (W) GENE

=> s L7 and microb?
L12          61 L7 AND MICROB?

=> s L12 and Corynebacterium
L13          1 L12 AND CORYNEBACTERIUM

=> dup rem L3
PROCESSING COMPLETED FOR L3
L14          26 DUP REM L3 (2 DUPLICATES REMOVED)

=> dup rem L4
PROCESSING COMPLETED FOR L4
L15          13 DUP REM L4 (0 DUPLICATES REMOVED)
```

```
=> dis L13 ibib kwic
```

```
L13 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2001 ACS
ACCESSION NUMBER:      1997:475788 CAPLUS
DOCUMENT NUMBER:       127:172154
TITLE:                 L-Lysine export from Corynebacterium
                        glutamicum. Physiological and molecular-biological
                        characterization of the carrier-mediated export of a
                        primary metabolite
AUTHOR(S):             Vrljic, Maria-Marina
CORPORATE SOURCE:       Inst. Biotechnologie, Forschungszentrum Julich
                        G.m.b.H., Juelich, D-52425, Germany
SOURCE:                Ber. Forschungszent. Juelich (1997), Juel-3349, 1-115
                        pp.
                        CODEN: FJBEE5; ISSN: 0366-0885
DOCUMENT TYPE:         Report
LANGUAGE:              German
```

```
TI L-Lysine export from Corynebacterium glutamicum. Physiological
and molecular-biological characterization of the carrier-mediated export
of a primary metabolite
AB The gene for the Lys-excretion carrier was isolated from C. glutamicum
and
the Lys export was analyzed physiol. A system was established which
induces the Lys excretion in dependence of Met. The mutant NA8, defect
in
Lys export, was isolated. The L-Lys export (LysE) gene encodes a
polypeptide of 236 amino acids with the potential to span the membrane 6
times and a mol. wt. of 2,5425 Da. With overexpressed LysE, L-Lys was
exported at a rate of 3.76 nmol/min/mg dry wt. which lead to a 10-fold
increased Lys excretion rate. The LysG (governing L-Lys export)
gene is localized immediately adjacent to LysE, but is
transcribed divergently. The deduced polypeptide (290 amino acids) has
a
helix-turn-helix motive at the aminotermminus. At the sequence level,
LysG
shows .ltoreq.35% identity to prokaryotic, autoregulatory transcriptional
```

regulators. LysG acts in trans and leads to a decrease of the Lys excretion by *C. glutamicum*. For the Lys-export defect mutant *C. glutamicum* NA8, the transition G1594.fwdarw.A1594 was shown which results in a stop-codon in the LysE gene. The resulting LysE polypeptide in *C. glutamicum* NA8 is shortened for 43 amino acids. The growth of a LysEG deletion mutant was abolished on a minimal medium in the presence of Lys-contg. dipeptides. The quantification of the intracellular L-Lys concns. revealed an accumulation of Lys .ltoreq.1,100 mM. The results suggest that the physiol. function of the Lys export carrier of *C. glutamicum* is to avoid extremely high intracellular Lys concns.

ST lysine excretion carrier **Corynebacterium** gene sequence; protein sequence **Corynebacterium** lysine excretion carrier

IT Amino acid transport (biological)  
(carrier-mediated, export; lysine export from **Corynebacterium** glutamicum, carrier-supported export of a primary metabolite)

IT Helix-turn-helix  
(gene lysG protein; lysine export from **Corynebacterium** glutamicum, carrier-supported export of a primary metabolite)

IT Proteins (specific proteins and subclasses)  
RL: BAC (Biological activity or effector, except adverse); BOC  
(Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(gene lysG, (governing lysine export); lysine export from **Corynebacterium** glutamicum, carrier-supported export of a primary metabolite)

IT Genes (**microbial**)  
RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(lysE; lysine export from **Corynebacterium** glutamicum, carrier-supported export of a primary metabolite)

IT Genes (**microbial**)  
RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(lysG (governing lysine export); lysine export from **Corynebacterium** glutamicum, carrier-supported export of a primary metabolite)

IT **Corynebacterium** glutamicum  
DNA sequences  
Protein sequences  
(lysine export from **Corynebacterium** glutamicum, carrier-supported export of a primary metabolite)

IT Amino acid transporters  
RL: BAC (Biological activity or effector, except adverse); BOC  
(Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(lysine-transporting, gene lysE; lysine export from **Corynebacterium** glutamicum, carrier-supported export of a primary metabolite)

IT 63-68-3, L-Methionine, biological studies  
RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)  
(induces lysine excretion; lysine export from **Corynebacterium** glutamicum, carrier-supported export of a primary metabolite)

IT 184922-77-8, GenBank X96471-derived protein GI 1729755  
RL: BAC (Biological activity or effector, except adverse); BOC  
(Biological occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)  
(lysine export from **Corynebacterium** glutamicum,

carrier-supported export of a primary metabolite)  
 IT 184922-76-7, GenBank X96471-derived protein GI 1729754 184922-78-9,  
 GenBank X96471-derived protein GI 1729756  
 RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological  
 study); OCCU (Occurrence)  
 (lysine export from **Corynebacterium** glutamicum,  
 carrier-supported export of a primary metabolite)  
 IT 56-87-1, L-Lysine, biological studies  
 RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
 (lysine export from **Corynebacterium** glutamicum,  
 carrier-supported export of a primary metabolite)  
 IT 184343-19-9, GenBank X96471  
 RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological  
 study); OCCU (Occurrence)  
 (nucleotide sequence; lysine export from **Corynebacterium**  
 glutamicum, carrier-supported export of a primary metabolite)

=> dis L14 1-26 ibib kwic

L14 ANSWER 1 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS  
 ACCESSION NUMBER: 2001:314459 BIOSIS  
 DOCUMENT NUMBER: PREV200100314459  
 TITLE: Effect of gluconic acid as a secondary carbon source on  
 non-growing L-lysine producers cells of  
**Corynebacterium** glutamicum. Purification and  
 properties of 6-phosphogluconate dehydrogenase.  
 AUTHOR(S): Bianchi, Daniella; Bertrand, Olivier; Haupt, Karsten;  
 Coello, Nereida (1)  
 CORPORATE SOURCE: (1) Instituto de Biologia Experimental, Universidad  
 Central  
 deVenezuela, Caracas, 1041-A: ncoello@uole.com.ve  
 Venezuela  
 SOURCE: Enzyme and Microbial Technology, (June 7, 2001) Vol. 28,  
 No. 9-10, pp. 754-759. print.  
 ISSN: 0141-0229.  
 DOCUMENT TYPE: Article  
 LANGUAGE: English  
 SUMMARY LANGUAGE: English  
 TI Effect of gluconic acid as a secondary carbon source on non-growing  
 L-lysine producers cells of **Corynebacterium** glutamicum.  
 Purification and properties of 6-phosphogluconate dehydrogenase.  
 AB We studied the production of L-lysine in **Corynebacterium**  
 glutamicum ATCC 21543 non growing cells obtained by nutrient limitation.  
 Statistical analysis revealed significant differences in the L-lysine  
 titers of. . .  
 IT . . .  
 Engineering; Methods and Techniques; Nutrition  
 IT Chemicals & Biochemicals  
 6-phosphogluconate dehydrogenase: amino acid sequence, analysis,  
 molecular properties, pH, purification; L-lysine: **microbial**  
**production**, yield; **amino acids**: analysis;  
 carbon sources; gluconic acid: secondary carbon source  
 ORGN . . .  
 Microorganisms; Irregular Nonsporing Gram-Positive Rods: Actinomycetes  
 and Related Organisms, Eubacteria, Bacteria, Microorganisms;  
 Microorganisms  
 ORGN Organism Name



Bacillus subtilis (Endospore-forming Gram-Positives);  
**Corynebacterium** glutamicum (Irregular Nonsporing Gram-Positive  
Rods): non-growing cells; Escherichia coli (Enterobacteriaceae);  
bacteria (Bacteria); microorganisms (Microorganisms)

ORGN Organism Superterms

Bacteria; Eubacteria; Microorganisms

L14 ANSWER 2 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER: 2001:421002 BIOSIS

DOCUMENT NUMBER: PREV200100421002

TITLE: L-glutamate fermentation and metabolic engineering:  
Studies

on the L-glutamate production mechanism in Coryneform  
bacteria.

AUTHOR(S): Nakamatsu, Tsuyoshi

SOURCE: Nippon Nogeikagaku Kaishi, (Jun., 2001) Vol. 75, No. 6,  
pp.

683-686. print.

ISSN: 0002-1407.

DOCUMENT TYPE: General Review

LANGUAGE: Japanese

SUMMARY LANGUAGE: English

IT Major Concepts

Biochemistry and Molecular Biophysics; Bioprocess Engineering;  
Metabolism

IT Chemicals & Biochemicals

**amino acids: large-scale microbial**

**production; glutamate: large-scale microbial**

**production; oxoglutarate dehydrogenase**

ORGN Super Taxa

Irregular Nonsporing Gram-Positive Rods: Actinomycetes and Related  
Organisms, Eubacteria, Bacteria, Microorganisms

ORGN Organism Name

**Corynebacterium** spp. (Irregular Nonsporing Gram-Positive  
Rods)

ORGN Organism Superterms

Bacteria; Eubacteria; Microorganisms

L14 ANSWER 3 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER: 2001:174925 BIOSIS

DOCUMENT NUMBER: PREV200100174925

TITLE: MALDI-TOF MS for quantification of substrates and products  
in cultivations of **Corynebacterium** glutamicum.

AUTHOR(S): Wittmann, Christoph (1); Heinzle, Elmar

CORPORATE SOURCE: (1) Biochemical Engineering Institute, Saarland  
University,

66041, Saarbruecken: c.wittmann@rz.uni-sb.de Germany

SOURCE: Biotechnology and Bioengineering, (March 20, 2001) Vol.  
72,

No. 6, pp. 642-647. print.

ISSN: 0006-3592.

DOCUMENT TYPE: Article

LANGUAGE: English

SUMMARY LANGUAGE: English

TI MALDI-TOF MS for quantification of substrates and products in  
cultivations

of **Corynebacterium** glutamicum.

IT Major Concepts

Biochemistry and Molecular Biophysics; Bioprocess Engineering; Methods  
and Techniques  
IT Chemicals & Biochemicals  
    **amino acids: microbial production**  
    , quantitative analysis; products: quantitative analysis; substrates:  
    quantitative analysis

L14 ANSWER 4 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:761605 CAPLUS

DOCUMENT NUMBER: 134:99608

TITLE: Development and use of miniaturized parallel  
experiment technology for bioprocess development

AUTHOR(S): Altenbach-Rehm, Jutta

CORPORATE SOURCE: Institut fur Biotechnologie, Julich, JUL-3782,  
Germany

SOURCE: Ber. Forschungszent. Juelich (2000), Juel-3782, a-f,  
i-iv, 1-233

CODEN: FJBEE5; ISSN: 0366-0885

DOCUMENT TYPE: Report

LANGUAGE: German

AB The fed-batch technique is nowadays the std. operation mode for high  
performance **microbial prodn.** processes. Shake flasks  
are widely used a simple bioreactors in batch process development.  
Because of uncontrolled changes in pH and reduced oxygen transfer rates  
parallel operated shake flasks can usually not be applied for microbial  
fed-batch process development. Due to the need to operate controlled  
stirred tank reactors the development of specific fermn. strategies is  
expensive and time consuming. To address these issues a miniature  
fed-batch technique was developed in cooperation with DASGIP mbH, Julich,  
and INFORS AG, Basel. Controlled batch and fed-batch fermns. can be  
performed in 16 parallel small scale bioreactors. The new technique  
allows the feeding of up to 4 different substrates and parallel pH  
control  
based on user-defined profiles. The feeding assembly is sterilized chem.  
using dimethylcarbonate. To overcome the limitations of shake flasks,  
small scale bubble columns were developed. Gas distribution is performed  
with a new type of sterile sparger. Transferring fed-batch fermns. from  
small scale bubble columns to a stirred tank reactor a scale up factor of  
20 was achieved. Escherichia coli K12 was chosen to test the new  
parallel  
bioreactor technique. Compared to shake flask fermns. the cell concn.  
was

50% higher due to efficient oxygen transfer. In fed-batch fermns. with  
pH-controlled substrate feeding up to 35 g/l DCW were achieved. For the  
1st time, a parallel optimization of feeding profiles in parallel small  
scale fed-batch expts. with successful scale-up to a lab. bioreactor was  
performed. Escherichia coli BL 21 (DE3) pLySS produces the recombinant  
enzyme GDP-.alpha.-D-mannose-pyrophosphorylase after induction with  
isopropyl-.beta.-D-thiogalactoside (IPTG). Single induction with 0,5 mM  
IPTG resulted in a low specific enzyme activity of 1,6 U/g DCW (dry cell  
wt.). For the optimization of enzyme expression a genetic algorithm was  
used. A final enzyme activity of 111 U/g DCW was achieved for optimal  
substrate and inducer feeding profiles. To demonstrate the advantages of  
this new parallel bioreactor technique different strains of industrial  
relevance were investigated. **Corynebacterium glutamicum**,  
Staphylococcus carnosus and Ashbya gossypii.

ST bioreactor bubble column fed batch fermn; GDP mannose pyrophosphorylase  
bubble column fed batch; Staphylococcus bubble column fed batch fermn;

isoleucine bubble column fed batch **Corynebacterium**; riboflavin  
bubble column fed batch Ashbya

IT **Amino acids**, biological studies  
RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
(amino acid consumption in riboflavin prodn. by Ashybya gossypii in  
parallel bubble columns with fed-batch technique)

IT **Corynebacterium** glutamicum  
(L-isoleucine prodn. by **Corynebacterium** glutamicum in  
parallel bubble columns with fed-batch technique)

IT 73-32-5P, L-Isoleucine, biological studies  
RL: BMF (Bioindustrial manufacture); BPR (Biological process); BIOL  
(Biological study); PREP (Preparation); PROC (Process)  
(L-isoleucine prodn. by **Corynebacterium** glutamicum in  
parallel bubble columns with fed-batch technique, amino acid  
consumption in riboflavin prodn. by Ashybya gossypii)

IT 61-90-5, L-Leucine, biological studies  
RL: BPR (Biological process); MFM (Metabolic formation); BIOL (Biological  
study); FORM (Formation, nonpreparative); PROC (Process)  
(L-isoleucine prodn. by **Corynebacterium** glutamicum in  
parallel bubble columns with fed-batch technique, amino acid  
consumption in riboflavin prodn. by Ashybya gossypii)

L14 ANSWER 5 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1999:244776 CAPLUS

DOCUMENT NUMBER: 130:266420

TITLE: Method for **microbial production** of  
**amino acids** of the aspartate and/or  
glutamate family and agents which can be used in said  
method

INVENTOR(S): Eikmanns, Bernd; Peters-Wendisch, Petra; Sahm,  
Hermann

PATENT ASSIGNEE(S): Forschungszentrum Julich G.m.b.H., Germany

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9918228	A2	19990415	WO 1998-EP6210	19980930
WO 9918228	A3	19990520		
W:	AU, BR, CA, CN, HU, ID, JP, KR, MX, RU, SK, US			
RW:	AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE			
DE 19831609	A1	19990415	DE 1998-19831609	19980714
AU 9911482	A1	19990427	AU 1999-11482	19980930
EP 1015621	A2	20000705	EP 1998-954301	19980930
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI			
BR 9813021	A	20000815	BR 1998-13021	19980930
PRIORITY APPLN. INFO.:			DE 1997-19743894 A	19971004
			DE 1998-19831609 A	19980714
			WO 1998-EP6210 W	19980930

TI Method for **microbial production** of **amino  
acids** of the aspartate and/or glutamate family and agents which  
can be used in said method

AB The invention relates to a method for **microbial prodn.** of **amino acids** of the aspartate and/or glutamate family in which the pyruvate carboxylase activity is increased by genetically changing the enzyme and/or the pyruvate carboxylase gene expression of a microorganism which produces the corresponding amino acid.

In addn., the invention relates to a pyruvate carboxylase gene and addnl. agents which can be used in the inventive method.

ST amino acid fermm **Corynebacterium** pyruvate carboxylase genetic engineering

IT **Corynebacterium** glutamicum  
Fermentation  
(**microbial prodn. of amino acids** of the aspartate and/or glutamate family and agents which can be used in said method)

IT **Amino acids**, preparation  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(**microbial prodn. of amino acids** of the aspartate and/or glutamate family and agents which can be used in said method)

IT Genetic engineering  
(**microbial prodn. of amino acids** of the aspartate and/or glutamate family and modification of **Corynebacterium** pyc gene in said method)

IT Genes (microbial)  
RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
(pyc; **microbial prodn. of amino acids** of the aspartate and/or glutamate family and modification of **Corynebacterium** pyc gene in said method)

IT 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, preparation 72-19-5P, L-Threonine, preparation 74-79-3P, L-Arginine, preparation 672-15-1P, L-Homoserine  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(**microbial prodn. of amino acids** of the aspartate and/or glutamate family and agents which can be used in said method)

IT 9014-19-1, Pyruvate carboxylase 204116-18-7 208541-81-5  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(**microbial prodn. of amino acids** of the aspartate and/or glutamate family and agents which can be used in said method)

L14 ANSWER 6 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER: 1999:175414 BIOSIS

DOCUMENT NUMBER: PREV199900175414

TITLE: Cloning of the transketolase gene and the effect of its dosage on aromatic amino acid production in **Corynebacterium** glutamicum.

AUTHOR(S): Ikeda, M. (1); Okamoto, K.; Katsumata, R.

CORPORATE SOURCE: (1) Technical Research Laboratories, Kyowa Hakko Kogyo Co.,

SOURCE: Ltd., Hofu, Yamaguchi, 747-8522 Japan  
Applied Microbiology and Biotechnology, (Feb., 1999) Vol. 51, No. 2, pp. 201-206.  
ISSN: 0175-7598.

DOCUMENT TYPE: Article

LANGUAGE: English

TI Cloning of the transketolase gene and the effect of its dosage on aromatic amino acid production in **Corynebacterium** glutamicum.

AB. . . enzyme of the non-oxidative pentose phosphate pathway. The effect of its overexpression on aromatic amino acid production was investigated in **Corynebacterium** glutamicum, a typical amino-acid-producing organism. For this purpose, the transketolase gene of the organism was cloned on the basis of. . . the presence of the gene in high copy numbers enabled tyrosine, phenylalanine and tryptophan producers to accumulate 5%-20% more aromatic **amino acids**. These results indicate that overexpressed transketolase activity operates to redirect the glycolytic intermediates toward the nonoxidative pentose phosphate pathway in. . .

IT Major Concepts  
Bioprocess Engineering; Enzymology (Biochemistry and Molecular Biophysics); Molecular Genetics (Biochemistry and Molecular Biophysics)

IT Chemicals & Biochemicals  
aromatic **amino acids: microbial production**; transketolase [EC 2.2.1.1]; **Corynebacterium** transketolase gene (Irregular Nonsporing Gram-Positive Rods)

L14 ANSWER 7 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1998:277651 CAPLUS

DOCUMENT NUMBER: 128:307587

TITLE: **Microbial production** of substances from aromatic metabolism

INVENTOR(S): Sprenger, Georg; Siewe, Ruth; Sahm, Hermann; Karutz, Martin; Sonke, Theodor

PATENT ASSIGNEE(S): Forschungszentrum Juelich G.m.b.H., Germany; Holland Sweetener Co. V.o.F.

SOURCE: Ger. Offen., 14 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
DE 19644566	A1	19980430	DE 1996-19644566	19961026
WO 9818936	A1	19980507	WO 1997-NL582	19971017
W:	AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, HU, ID, IL, IS, JP, KP, KR, LC, LK, LR, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
RW:	GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG			
AU 9747277	A1	19980522	AU 1997-47277	19971017
EP 934418	A1	19990811	EP 1997-909748	19971017
R:	AT, BE, CH, DE, DK, ES, FR, GB, IT, LI, NL, SE, PT, FI			
CN 1241214	A	20000112	CN 1997-180908	19971017
JP 2001506486	T2	20010522	JP 1998-520318	19971017
PRIORITY APPLN. INFO.:			DE 1996-19644566 A	19961026
			WO 1997-NL582 W	19971017

TI **Microbial production** of substances from aromatic metabolism

IT Transport proteins  
 RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU (Biological use, unclassified); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)  
 (gene glf glucose facilitator protein, of *Zymomonas mobilis*;  
**microbial prodn.** of substances from arom. metab.)

IT Genes (microbial)  
 RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
 (glf, for glucose facilitator protein of *Zymomonas mobilis*;  
**microbial prodn.** of substances from arom. metab.)

IT Genes (microbial)  
 RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
 (glk, for glucokinase of *Zymomonas mobilis*; **microbial prodn.** of substances from arom. metab.)

IT Pentose phosphate pathway  
 (intermediates of, in amino acid manuf.; **microbial prodn.** of substances from arom. metab.)

IT *Bacillus* (bacterium genus)  
*Brevibacterium*  
*Corynebacterium*  
*Escherichia*  
*Escherichia coli*  
 Fermentation  
 Microorganism  
 Molecular cloning  
*Serratia*  
 (**microbial prodn.** of substances from arom. metab.)

IT Transport proteins  
 RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU (Biological use, unclassified); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)  
 (**microbial prodn.** of substances from arom. metab.)

IT **Amino acids**, preparation  
 RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
 (**microbial prodn.** of substances from arom. metab.)

IT Plasmids  
 (pZ4557tal; **microbial prodn.** of substances from arom. metab.)

IT Plasmids  
 (pZ4557tkk; **microbial prodn.** of substances from arom. metab.)

IT Plasmids  
 (pZ4557tkktal; **microbial prodn.** of substances from arom. metab.)

IT Genes (microbial)  
 RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
 (talB; **microbial prodn.** of substances from arom. metab.)

IT Genes (microbial)  
 RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
 (tkkA; **microbial prodn.** of substances from arom. metab.)

metab.)

IT 9001-36-9P, Glucokinase  
 RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU (Biological use, unclassified); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)  
 (gene glk, of *Zymomonas mobilis*; **microbial prodn.** of substances from arom. metab.)

IT 585-18-2, Erythrose-4-phosphate  
 RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses)  
 (in amino acid manuf.; **microbial prodn.** of substances from arom. metab.)

IT 9014-46-4P, Transaldolase 9014-48-6P, Transketolase  
 RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU (Biological use, unclassified); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); USES (Uses)  
 (**microbial prodn.** of substances from arom. metab.)

IT 63-91-2P, L-Phenylalanine, preparation  
 RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
 (**microbial prodn.** of substances from arom. metab.)

L14 ANSWER 8 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1995:674124 CAPLUS  
 DOCUMENT NUMBER: 123:54314  
 TITLE: Enhancement of reduced NADP production for enhanced **microbial production** of biochemicals  
 INVENTOR(S): Kojima, Hiroyuki; Totsuka, Kazuhiko  
 PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan  
 SOURCE: PCT Int. Appl., 32 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: Japanese  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9511985	A1	19950504	WO 1994-JP1791	19941026
W: AU, BR, CA, CN, CZ, HU, JP, KR, PL, RU, SK, US, VN				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
CA 2175042	AA	19950504	CA 1994-2175042	19941026
AU 9480026	A1	19950522	AU 1994-80026	19941026
AU 687458	B2	19980226		
EP 733712	A1	19960925	EP 1994-931158	19941026
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT,				
SE				
BR 9407907	A	19961126	BR 1994-7907	19941026
HU 74840	A2	19970228	HU 1996-1085	19941026
ZA 9503350	A	19961025	ZA 1995-3350	19950425
US 5830716	A	19981103	US 1996-619521	19960429
CN 1139956	A	19970108	CN 1994-194707	19961026
PRIORITY APPLN. INFO.:			JP 1993-270828	19931028
			WO 1994-JP1791	19941026

TI Enhancement of reduced NADP production for enhanced **microbial production** of biochemicals

AB The productivity of such substances as L-amino acids, antibiotics, vitamins, growth factors and physiol. active substances in

the fermn. using a microorganism is improved by improving the productivity of reduced NADP in the cells of the microorganisms. Construction of pMW::THY contg. the Escherichia coli transhydrogenase gene, and introduction of the plasmid into the L-threonine-producing Escherichia coli B-3996 were shown. The recombinant E. coli B-3996 produced L-threonine .apprx.10% higher than did the parental strain.

IT **Corynebacterium glutamicum**  
Escherichia coli  
Fermentation  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT **Amino acids**, preparation  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT Plasmid and Episome  
(pHSG::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT Plasmid and Episome  
(pMW::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT Plasmid and Episome  
(pSU::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT 9014-18-0, Nicotinamide nucleotide transhydrogenase  
RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, preparation 61-90-5P, L-Leucine, preparation 63-91-2P, L-Phenylalanine, preparation 72-18-4P, L-Valine, preparation 72-19-5P, L-Threonine, preparation 73-32-5P, L-Isoleucine, preparation  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT 53-59-8P, NADP  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(reduced; enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

L14 ANSWER 9 OF 26 SCISEARCH COPYRIGHT 2001 ISI (R)  
ACCESSION NUMBER: 95:184306 SCISEARCH  
THE GENUINE ARTICLE: QK574  
TITLE: METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM  
**CORYNEBACTERIUM-GLUTAMICUM**  
AUTHOR: SAHM H (Reprint); EGDELING L; EIKMANN B; KRAMER R  
CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL, D-52425 JULICH, GERMANY (Reprint)  
COUNTRY OF AUTHOR: GERMANY  
SOURCE: FEMS MICROBIOLOGY REVIEWS, (FEB 1995) Vol. 16, No. 2-3, pp. 243-252.  
ISSN: 0168-6445.  
DOCUMENT TYPE: Article; Journal



FILE SEGMENT: LIFE  
LANGUAGE: ENGLISH  
REFERENCE COUNT: 36

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

TI METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM **CORYNEBACTERIUM**  
-GLUTAMICUM

AB The Gram-positive bacterium **Corynebacterium** glutamicum is  
used for the industrial production of **amino acids**,  
e.g. of L-glutamate and L-lysine. In the last 10 years, genetic  
engineering and amplification of relevant structural genes have become.

ST Author Keywords: **CORYNEBACTERIUM** GLUTAMICUM; AMINO ACID  
PRODUCTION; METABOLIC DESIGN; L-LYSINE; L-THREONINE; L-ISOLEUCINE

STP KeyWords Plus (R): L-THREONINE; L-LYSINE; BREVIBACTERIUM-LACTOFERMENTUM;  
HOMOSERINE DEHYDROGENASE; **MICROBIAL PRODUCTION**;  
RESISTANT MUTANTS; SPLIT PATHWAY; BIOSYNTHESIS; ISOLEUCINE; GENES

L14 ANSWER 10 OF 26 SCISEARCH COPYRIGHT 2001 ISI (R)

ACCESSION NUMBER: 96:7187 SCISEARCH

THE GENUINE ARTICLE: TJ545

TITLE: METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM  
**CORYNEBACTERIUM**-GLUTAMICUM

AUTHOR: SAHM H (Reprint)

CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL,  
D-52425 JULICH, GERMANY (Reprint)

COUNTRY OF AUTHOR: GERMANY

SOURCE: FOLIA MICROBIOLOGICA, (1995) Vol. 40, No. 1, pp. 23-30.  
ISSN: 0015-5632.

DOCUMENT TYPE: Article; Journal

FILE SEGMENT: LIFE; AGRI

LANGUAGE: ENGLISH

REFERENCE COUNT: 26

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

TI METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM  
**CORYNEBACTERIUM**-GLUTAMICUM

AB The Gram-positive bacterium **Corynebacterium** glutamicum is  
used for the industrial production of **amino acids**,  
e.g. of L-glutamate and L-lysine. By cloning and expressing the various  
genes of the L-lysine pathway in *C. glutamicum* we. . .

STP KeyWords Plus (R): L-THREONINE; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE  
DEHYDROGENASE; **MICROBIAL PRODUCTION**; LYSINE  
BIOSYNTHESIS; RESISTANT MUTANTS; SPLIT PATHWAY; GENES; AMPLIFICATION;  
FERMENTATION

L14 ANSWER 11 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1985:436156 CAPLUS

DOCUMENT NUMBER: 103:36156

TITLE: Optimization of amino acid production by automatic  
self-tuning digital control of redox potential

AUTHOR(S): Radjai, Mohammad K.; Hatch, Randolph T.; Cadman,  
Theodore W.

CORPORATE SOURCE: Dep. Chem. Nucl. Eng., Univ. Maryland, College Park,  
MD, 20742, USA

SOURCE: Biotechnol. Bioeng. Symp. (1984), 14 (Symp.  
Biotechnol.

Fuels Chem., 6th), 657-79

CODEN: BIBSBR; ISSN: 0572-6565

DOCUMENT TYPE: Journal  
LANGUAGE: English

AB The **microbial prodn.** of homoserine [672-15-1], lysine [56-87-1], and valine [72-18-4] by an auxotrophic mutant of **Corynebacterium glutamicum** was investigated in a 16-L batch fermentor. Closed-loop digital control of redox potential was

implemented

using proportional-integral (PI) control of agitation rates. Due to the nonlinearity of the system, the PI controller parameters had to be changed

during the course of the fermns. An automatic, self-tuning algorithm was developed for stable control of redox potential. This permitted exptl. optimization of total and selective amino acid prodn. Total amino acid yields of 35% from glucose were achieved compared to 23% reported in the literature for the same fermn.

IT **Corynebacterium glutamicum**

(amino acid manuf. with, optimization and redox potential control in)

IT **Amino acids**, preparation

RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP (Preparation)  
(manuf. of, by fermn.)

L14 ANSWER 12 OF 26 CAPLUS COPYRIGHT 2001 ACS DUPLICATE 1

ACCESSION NUMBER: 1979:20814 CAPLUS

DOCUMENT NUMBER: 90:20814

TITLE: **Microbial production** of essential  
**amino acids** with

**Corynebacterium glutamicum** mutants

AUTHOR(S): Nakayama, Kiyoshi; Araki, Kazumi; Kase, Hiroshi

CORPORATE SOURCE: Tokyo Res. Lab., Kyowa Hakko Kogyo Co., Ltd.,  
Machida,

Japan

SOURCE: Adv. Exp. Med. Biol. (1978), 105(Nutr. Improv. Food  
Feed Proteins), 649-61

CODEN: AEMBAP; ISSN: 0065-2598

DOCUMENT TYPE: Journal

LANGUAGE: English

TI **Microbial production** of essential **amino**  
**acids** with **Corynebacterium glutamicum** mutants

AB **Amino acids** produced by microbial processes are generally L-forms. The stereospecificity of the **amino acids** produced by fermn. makes the process advantageous compared with synthetic processes. Microorganisms employed in microbial processes for amino acid prodn. are divided into 4 classes: wild-type, auxotrophic mutant, regulatory mutant, and auxotrophic regulatory mutant. Using such mutants of **Corynebacterium glutamicum**, all the essential **amino acids** but L-methionine are now being produced by direct fermn. from cheap C sources such as carbohydrate materials or acetic acid.

ST amino acid manuf **Corynebacterium**

IT **Corynebacterium glutamicum**  
(amino acid manuf. by)

IT Fermentation  
(**amino acids**, by **Corynebacterium**  
glutamicum)

IT **Amino acids**, preparation

RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP (Preparation)

(manuf. of, from carbohydrates by **Corynebacterium** glutamicum)

L14 ANSWER 13 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1976:521806 CAPLUS

DOCUMENT NUMBER: 85:121806

TITLE: **Microbial production** of amino acid

INVENTOR(S): Tsuchida, Takayasu; Yoshihara, Yasuhiko; Kubota, Koji;

Hirose, Yoshio

PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan

SOURCE: Japan. Kokai, 5 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	---	----	-----	-----
	JP 51061690	A2	19760528	JP 1974-134879	19741122
TI	<b>Microbial production</b> of amino acid				
ST	amino acid manuf Brevibacterium; <b>Corynebacterium</b> amino acid manuf				
IT	Brevibacterium				
	<b>Corynebacterium</b>				
	(amino acid manuf. by)				
IT	Fermentation				
	(amino acids, by <b>Corynebacterium</b> or Brevibacterium)				
IT	56-45-1P, preparation 73-22-3P, preparation				
	RL: PREP (Preparation)				
	(by fermn., with <b>Corynebacterium</b> )				

L14 ANSWER 14 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1975:529947 CAPLUS

DOCUMENT NUMBER: 83:129947

TITLE: **Microbial production** of amino acids. VI. Formation of L-amino acids from DL-.alpha.-hydroxycarboxylic acids

AUTHOR(S): Matsushima, Hirochika; Murata, Keijiro; Mase, Yasuo

CORPORATE SOURCE: Ferment. Res. Lab., Sankyo Co., Ltd., Tanashi, Japan

SOURCE: Hakko Kogaku Zasshi (1975), 53(7), 443-9

CODEN: HKZAA2

DOCUMENT TYPE: Journal

LANGUAGE: Japanese

TI **Microbial production** of amino acids

. VI. Formation of L-amino acids from

DL-.alpha.-hydroxycarboxylic acids

AB Formation of L-amino acids from DL-.alpha.-

hydroxycarboxylic acids was studied. L-.alpha.-aminobutyric acid [1492-24-6] was formed in a medium contg. DL-.alpha.-hydroxybutyric acid [600-15-7] by various bacteria belonging to Aerobacter, Bacillus, **Corynebacterium**, Escherichia, Flavobacterium, Micrococcus, Proteus, Pseudomonas, Sarcina, Staphylococcus, and other genera. A. cloacae IAM 1221 was cultured in a medium contg. DL-.alpha.-bromobutyric acid [2385-70-8] (hydrolyzed to hydroxybutyric acid). L-.alpha.-aminobutyric acid was isolated from the culture broth and

identified by thin-layer chromatog., elementary anal., and by its specific rotation and IR spectrum. Formation of valine [72-18-4], leucine [61-90-5], or phenylalanine [63-91-2] from DL-.alpha.-hydroxycarboxylic acids by *Brevibacterium roseum* ATCC 13825 was studied. Yields (mole) from the cultures were 84.22, 95.7, and 47.7%, resp. An amino-group donor (glutamic acid) was needed besides the bacterial cells and DL-.alpha.-hydroxycarboxylic acid for the enzymic formation of **amino acids**.

L14 ANSWER 15 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1975:137747 CAPLUS

DOCUMENT NUMBER: 82:137747

TITLE: **Microbial production of amino acids**

INVENTOR(S): Kubota, Koji; Yoshihara, Yasuhiko; Okada, Hiroshi

PATENT ASSIGNEE(S): Ajinomoto Co., Inc.

SOURCE: Japan. Kokai, 5 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 49109585	A2	19741018	JP 1973-24049	19730228
JP 51038796	B4	19761023		

TI **Microbial production of amino acids**

AB **Amino acids** were produced by a microbe cultured in a propionic acid medium. Thus, *Brevibacterium flavum* ATCC 14,067, *Micrococcus glutamicus* ATCC 13,032, ***Corynebacterium*** *acetoacidophilum* ATCC 13,870, *Microbacterium ammoniaphilum* ATCC 15,354, and *B. flavum* FERM-P 1684 were cultured with shaking at 31.degree. for 48 hr in a medium (pH 7.5) contg. propionic acid 2, (NH<sub>4</sub>)<sub>2</sub>SO<sub>4</sub> 1, KH<sub>2</sub>PO<sub>4</sub> 0.1, MgSO<sub>4</sub>.cntdot.7H<sub>2</sub>O 0.04, NaCl 0.1, and soybean protein hydrolysate (total

N = 7%) 0.2% plus biotin 2 and thiamine.cntdot.HCl 200 .mu.g/l. Prodn. of L-glutamic acid by each organism was 4.3, 4.2, 3.9, 4.0, and 2.5 mg/ml, resp. *B. flavum* FERM-P 1684 also produced N-acetylglutamine at 0.4 mg/ml.

IT ***Corynebacterium* acetoacidophilum**  
(glutamic acid manuf. by, from propionic acid)

L14 ANSWER 16 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1973:56392 CAPLUS

DOCUMENT NUMBER: 78:56392

TITLE: **Microbial production of amino acids** from aromatic compounds.  
I. Screening of aromatic compound-assimilating bacteria

AUTHOR(S): Yamamoto, Masao; Nishida, Hiroshi; Inui, Taiji; Ozaki,

Asaichiro

CORPORATE SOURCE: Cent. Res. Lab., Sanraku-Ocean Co., Ltd., Fujisawa, Japan

SOURCE: Hakko Kogaku Zasshi (1972), 50(12), 868-75

CODEN: HKZAA2

DOCUMENT TYPE: Journal

LANGUAGE: English

TI **Microbial production of amino acids**

from aromatic compounds. I. Screening of aromatic compound-assimilating bacteria

AB In an attempt to produce **amino acids** from aromatic compds. by fermn., bacterial stock cultures in this lab. were examd. for their assimilability of benzoate and salicylate; 96 strains from 97 glutamate-producing cultures assimilated benzoic acid. Then, 10 type-strains of the glutamate-producing strains were tested for their assimilability of 40 aromatic compds. 16 of the compds. were assimilated. These were benzoic acid, m-hydroxybenzoic acid, p-hydroxybenzoic acid, 2,4-dihydroxybenzoic acid, 2,5-dihydroxybenzoic acid, 3,4-dihydroxybenzoic acid, 3,5-dihydroxybenzoic acid, phenylacetic acid, phenylpyruvic acid, .beta.-phenylpropionic acid, cinnamic acid, benzal dehyde, benzyl alc., phenol, catechol, and resorcinol. A sizable amt. of L-glutamic acid

was produced from the assimilated compds. by these glutamate-producing bacteria, benzoate, esp., serving as the best substrate.

IT Brevibacterium

Brevibacterium lactofermentum

**Corynebacterium** acetoglutamicum

Microbacterium ammoniaphilum

Micrococcus glutamicus

(glutamic acid formation by, from arom. compds.)

L14 ANSWER 17 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1971:84222 CAPLUS

DOCUMENT NUMBER: 74:84222

TITLE: Utilization of hydrocarbons by microorganisms. XXI.

Biochemical studies of **microbial**

**production** of .alpha.-ketoglutarate,

L-glutamate, and DL-alanine from hydrocarbons

AUTHOR(S): Imada, Yukio; Yamada, Koichi

CORPORATE SOURCE: Fac. Agric., Univ. Tokyo, Tokyo, Japan

SOURCE: Agr. Biol. Chem. (1971), 35(1), 18-26

CODEN: ABCHA6

DOCUMENT TYPE: Journal

LANGUAGE: English

TI Utilization of hydrocarbons by microorganisms. XXI. Biochemical studies of **microbial production** of .alpha.-ketoglutarate, L-glutamate, and DL-alanine from hydrocarbons

AB Strain S10B1 of **Corynebacterium** hydrocarboclastus produced .alpha.-ketoglutaric acid (I), L-glutamate, and DL-alanine from n-alkanes in a thiamine (II)-limited medium supplemented with Fe<sup>2+</sup>. The replacement of hydrocarbon substrate by sugars such as glucose not only decreased the yields, but also reversed the order of the yields among the 3 products. This phenomenon was explained by a metabolic pathway in relation to the role of II. Slow O<sub>2</sub> uptake in the presence of pyruvate and I by II-deficient cells supported the presumption that II limitation resulted

in deficiency of a cofactor in the enzymic oxidn. of pyruvate and I. Activities of terminal enzymes in the synthesis of L-glutamate and DL-alanine

were detd. and discussed. Three intermediates were detected in the culture broth.

ST **Corynebacterium** ketoglutarate prodn; ketoglutarate prodn

**Corynebacterium**; glutamate prodn **Corynebacterium**;  
alanine prodn **Corynebacterium**; thiamine **Corynebacterium**  
; hydrocarbons utilization bacteria; bacteria hydrocarbons utilization

IT **Corynebacterium**  
(hydrocarboclastus, amino acids formation by, from  
hydrocarbons)

IT 59-43-8, biological studies  
RL: BIOL (Biological study)  
(amino acids formation from paraffins by  
**Corynebacterium** hydrocarboclastus in response to)

L14 ANSWER 18 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1970:475660 CAPLUS

DOCUMENT NUMBER: 73:75660

TITLE: **Microbial production** of L-glutamic  
acid

PATENT ASSIGNEE(S): Asahi Chemical Industry Co., Ltd.

SOURCE: Fr. Demande, 11 pp.

CODEN: FRXXBL

DOCUMENT TYPE: Patent

LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
-----	----	-----	-----	-----
FR 2009795		19700206		
PRIORITY APPLN. INFO.:		JP		19680531
TI <b>Microbial production</b> of L-glutamic acid				
AB L-Glutamic acid (I) is prepd. by aerobic cultivation of <b>Corynebacterium</b> or Brevibacterium in liq. media contg. C1-3 alcs. as C source and penicillin. Thus, B. vitalumen var propanolophilum ATCC 21391 was grown in a medium contg. PrOH 50, corn steep liquor 4, KH <sub>2</sub> PO <sub>4</sub> 2, MgSO <sub>4</sub> ·7H <sub>2</sub> O 0.5, Fe <sup>2+</sup> 0.01, Mn <sup>2+</sup> 0.01, urea 4 g/l., with the addn. of 100 .mu.g biotin and penicillin G (K salt) 10 units/l., at 32.degree. and pH 6.5-8.0 with shaking for 96 hr to give 23.1 g I/l. (46.2% based on PrOH). PrOH and penicillin were added in portions during the fermentation. Without penicillin addn., the yield was 6.4% I.				
ST Brevibacterium glutamate prodn; glutamate prodn Brevibacterium; <b>amino acids Corynebacterium</b> ; <b>Corynebacterium amino acids</b> ; penicillin bacteria glutamate				
IT <b>Corynebacterium</b> (melassecola and petrophylum, glutamic acid manuf. by)				

L14 ANSWER 19 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1970:508239 CAPLUS

DOCUMENT NUMBER: 73:108239

TITLE: **Microbial production** of  
L-threonine

INVENTOR(S): Nakayama, Kiyoshi; Kase, Hiroshi

PATENT ASSIGNEE(S): Kyowa Fermentation Industry Co. Ltd.

SOURCE: Ger. Offen., 22 pp.

CODEN: GWXXBX

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	---	-----	-----	-----
	DE 1817666	A	19700827	DE 1968-1817666	19681224

TI **Microbial production** of L-threonine

AB Various microorganisms, e.g. Aerobacter [Enterobacter] aerogenes, Serratia marcescens, or Arthrobacter paraffineus, cultured for producing L-threonine required 2 or 3 of the **amino acids** isoleucine, methionine, lysine, or diaminopimelic acid. The microorganisms were cultured aerobically in an aq. medium contg. the optimal (or less) amts. of the required **amino acids**. Thus, E. aerogenes NM-IS-5 (ATCC 21,215) was cultured 96 hr at 30.degree. in medium contg. glucose 5, (NH<sub>4</sub>)<sub>2</sub>SO<sub>4</sub> 1.4, KH<sub>2</sub>PO<sub>4</sub> 0.05, K<sub>2</sub>HPO<sub>4</sub> 0.05, MgSO<sub>4</sub>.7H<sub>2</sub>O 0.025, FeSO<sub>4</sub>.7H<sub>2</sub>O 0.001, MnSO<sub>4</sub>.4H<sub>2</sub>O 0.001, and CaCO<sub>3</sub> 2% and isoleucine 50, methionine 100, and diaminopimelic acid 200 mg/l. to give 7.8 g L-threonine/l.

ST **microbial prodn** threonine; threonine **microbial prodn**; Aerobacter threonine fermn; amino acid prodn fermn

IT **Corynebacterium**  
(glutamicum, threonine manuf. by)

L14 ANSWER 20 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1970:401150 CAPLUS

DOCUMENT NUMBER: 73:1150

TITLE: **Microbial production** of L-threonine. II. Production by .alpha.-amino-.beta.-hydroxyvaleric acid resistant mutants of glutamate producing bacteria

AUTHOR(S): Shio, Isamu; Nakamori, Shigeru

CORPORATE SOURCE: Cent. Res. Lab., Ajinomoto Co., Inc., Kawasaki, Japan

SOURCE: Agr. Biol. Chem. (1970), 34(3), 448-56

CODEN: ABCHA6

DOCUMENT TYPE: Journal

LANGUAGE: English

TI **Microbial production** of L-threonine. II. Production by .alpha.-amino-.beta.-hydroxyvaleric acid resistant mutants of glutamate producing bacteria

AB A mutant strain of Brevibacterium flavum was able to grow in a medium contg. 5 mg DL-threo-.alpha.-amino-.beta.-hydroxyvaleric acid (AHV)/ml; 1 mg AHV/ml inhibited the growth of the parental strain by >90%. Further treatment of the AHV-resistant strain with the mutagen, N-methyl-N'-nitro-N-nitrosoguanidine, produced a bacterial strain that was able to grown on 8 mg AHV/ml; this mutant produced 13.5 g L-threonine/l., an amt. 30% more than that produced by the parental strain. A similarly derived mutant of **Corynebacterium** acetoacidophilum resistant to AHV produced 6.1 g threonine/l. Other **amino acids** biosynthesized by the bacteria were discussed in relation to the regulation of threonine synthesis.

ST threonine prodn bacterial; **corynebacterium** threonine prodn; Brevibacterium threonine prodn; mutations bacteria threonine; bacteria mutations threonine; aminohydroxyvalerate bacteria

IT **Corynebacterium**  
(acetoacidophilum, tryptophan formation from aminohydroxyvaleric acid

by)

L14 ANSWER 21 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS  
ACCESSION NUMBER: 1970:123860 BIOSIS  
DOCUMENT NUMBER: BA51:33860  
TITLE: **MICROBIAL PRODUCTION OF AMINO  
-ACIDS FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID  
PRODUCTION BY CORYNEBACTERIUM-HYDROCARBOCLASTUS-R-  
7.**

AUTHOR(S): SHIIO I; UCHIO R  
SOURCE: AMINO ACID NUCLEIC ACID, (1969) (19), 88-96.  
CODEN: HATAA4. ISSN: 0517-6174.

FILE SEGMENT: BA; OLD  
LANGUAGE: Unavailable

TI **MICROBIAL PRODUCTION OF AMINO-ACIDS  
FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID PRODUCTION BY  
CORYNEBACTERIUM-HYDROCARBOCLASTUS-R-7.**

L14 ANSWER 22 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS DUPLICATE 2  
ACCESSION NUMBER: 1970:106213 BIOSIS  
DOCUMENT NUMBER: BA51:16213  
TITLE: **MICROBIAL PRODUCTION OF AMINO  
-ACIDS FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID  
PRODUCTION BY CORYNEBACTERIUM-HYDROCARBOCLASTUS  
R-7.**

AUTHOR(S): SHIIO I; UCHIO R  
SOURCE: J GEN APPL MICROBIOL, (1969) 15 (1), 65-84.  
CODEN: JGAMA9. ISSN: 0022-1260.

FILE SEGMENT: BA; OLD  
LANGUAGE: Unavailable

TI **MICROBIAL PRODUCTION OF AMINO-ACIDS  
FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID PRODUCTION BY  
CORYNEBACTERIUM-HYDROCARBOCLASTUS R-7.**

L14 ANSWER 23 OF 26 CAPLUS COPYRIGHT 2001 ACS  
ACCESSION NUMBER: 1967:514494 CAPLUS  
DOCUMENT NUMBER: 67:114494  
TITLE: **Microbial production of  
amino acids from hydrocarbons. III.  
L-Ornithine production by an arginine auxotrophic  
mutant of Corynebacterium hydrocarboclastus**

AUTHOR(S): Ishu, Ryosuke; Ishii, Ryosuke; Shiio, Isamu  
CORPORATE SOURCE: Ajinomoto Co., Inc., Kawasaki, Japan  
SOURCE: J. Gen. Appl. Microbiol. (1967), 13(3), 3303-12  
CODEN: JGAMA9

DOCUMENT TYPE: Journal  
LANGUAGE: English

TI **Microbial production of amino acids  
from hydrocarbons. III. L-Ornithine production by an arginine  
auxotrophic mutant of Corynebacterium hydrocarboclastus**  
AB cf. CA 67: 89718u. The arginine auxotrophic mutant strain RN-362 of C.  
hydrocarboclastus R-7 was used to study L-ornithine production from  
hydrocarbons, in a fermentation medium contg. various n-alkanes.  
L-Ornithine production required L-arginine at the optimum level of  
0.5-1.0  
g./l. of medium; an excess inhibited the biosynthesis of L-ornithine.  
(NH<sub>4</sub>)<sub>2</sub>HPO<sub>4</sub> was the best source of N and, at 2% in a neutral to slightly  
acidic pH, gave the highest level of L-ornithine production and cell



growth; NH<sub>4</sub>OAc, KNO<sub>3</sub>, and (NH<sub>4</sub>)<sub>2</sub>CO<sub>3</sub> proved less suitable because of a drop in pH along with the accumulation of a large amt. of .alpha.-ketoglutaric acid, pyruvic acid, and proline in the growth medium. Of 17 C sources, n-tetradecane best supported cell growth and L-ornithine production and the other C<sub>13</sub>-C<sub>17</sub> n-alkanes did so moderately, while kerosene and light oil produced good cell growth but only a small amt. of L-ornithine.

Addn. of 3 g. yeast ext. and 0.5 g. L-arginine-HCl to 1 l. of medium enhanced L-ornithine production. A similar effect was achieved by replacing the yeast ext. with various **amino acids** at 0.01% in the medium. L-Methionine was most effective for the production of L-ornithine, while L-lysine, L-cysteine, L-cystine, L-histidine, and L-phenylalanine were less so, in decreasing order. **Amino acids** enhance L-ornithine production by stimulating hydrocarbon oxidn. and cell growth.

ST HYDROCARBONS USE BACTERIA; BACTERIA HYDROCARBONS USE; ALKANES USE BACTERIA; **AMINO ACIDS** PRODN HYDROCARBONS; ORNITHINE PRODN HYDROCARBONS; PARAFFINS UTILIZATION BACTERIA

IT **Corynebacterium**

(hydrocarboclastus, ornithine formation from hydrocarbons by)

IT Hydrocarbons, biological studies

RL: BIOL (Biological study)

(ornithine formation from, by **Corynebacterium** hydrocarboclastus)

IT 70-26-8

RL: FORM (Formation, nonpreparative)

(formation of, from hydrocarbons by **Corynebacterium** hydrocarboclastus)

L14 ANSWER 24 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1967:489718 CAPLUS

DOCUMENT NUMBER: 67:89718

TITLE: **Microbial production of amino acids** from hydrocarbons. II.

Isolation of good hydrocarbon utilizers and amino acid production by their auxotrophs

AUTHOR(S): Ishii, Ryosuke; Otsuka, Shinichiro; Shio, Isamu  
CORPORATE SOURCE: Central Res. Labs., Ajinomoto Co., Inc., Kawasaki, Japan

SOURCE: J. Gen. Appl. Microbiol. (1967), 13(2), 217-25  
CODEN: JGAMA9

DOCUMENT TYPE: Journal

LANGUAGE: English

TI **Microbial production of amino acids**  
from hydrocarbons. II. Isolation of good hydrocarbon utilizers and amino acid

production by their auxotrophs

AB cf. CA 59: 14313h. Nine microorganisms, which showed good growth on long-chain aliphatic hydrocarbons, were isolated by an enrichment culture method, followed by a single colony isolation technique. They included 5 strains of *Alcaligenes marshallii*, 2 strains of **Corynebacterium** hydrocarboclastus, and 2 strains of yeast. Various auxotrophic mutants were derived from these microorganisms. The mutants accumulated the following **amino acids** from aliphatic hydrocarbons; L-ornithine, L-valine, L-glutamic acid, L-leucine, L-tyrosine, L-alanine, L-proline, L-aspartic acid, and L-lysine.

ST BACTERIA AMINO ACID PRODN; AMINO ACID PRODN BACTERIA; HYDROCARBONS

**AMINO ACIDS; ALIPHATICS BACTERIA METAB**

IT **Corynebacterium**  
(hydrocarboclastus, amino acid fermentation of hydrocarbons by)  
IT **Amino acids, preparation**  
RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP  
(Preparation)  
(manuf. of, by fermentation of hydrocarbons)

L14 ANSWER 25 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1966:22870 CAPLUS

DOCUMENT NUMBER: 64:22870

ORIGINAL REFERENCE NO.: 64:4230g-h,4231a

TITLE: **Microbial production of nucleotides**

INVENTOR(S): Masuo, Eitaro; Okabayashi, Tadashi

PATENT ASSIGNEE(S): Shionogi & Co., Ltd.

SOURCE: 10 pp.

DOCUMENT TYPE: Patent

LANGUAGE: Unavailable

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 40010957		19650601	JP	19591214

TI **Microbial production of nucleotides**

AB Some bacteria strains of high nucleotide-forming activity were detected based on the results of the test developed by the authors, and compns. of media for promoting accumulation of nucleotides were also investigated. To evaluate the nucleotide-forming activity of bacteria, cells of nonexacting purine (I) auxotrophic mutant B 96 of Escherichia coli were mixed into the synthetic medium contg. no I for testing strains. The activity of nucleotide accumulation of the strains increased as the growth

of the mutant increased. By this procedure, the following strains were found to be suitable for nucleotide production: Bacillus subtilis IFO 3061, B. firmus IFO 3330, B. circulans IFO 3342, B. megaterium IFO 3003, Alcaligenes viscosus AN-14, A. metalcaligenes 1021, Serratia marcescens 1008, S. plymuthica IFO 3055, Bacterium ketoglutaricum 1041, and new species of Brevibacterium and **Corynebacterium**. For promoting nucleotide production with these strains, **amino acids**, esp. L-glutamic acid (II), are necessary in the medium. Proteins or peptides contg. II are also effective for the strains having sufficient protease. Sufficient content of PO43- at pH 5.0-7.5 is also necessary

for the medium. By cultivation under these conditions, AMP, CDP, UMP, and UDP are obtained.

L14 ANSWER 26 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1963:476777 CAPLUS

DOCUMENT NUMBER: 59:76777

ORIGINAL REFERENCE NO.: 59:14313h,14314a

TITLE: **Microbial production of amino acids from hydrocarbons. I.**  
Preliminary screening of glutamic acid-producing bacteria

AUTHOR(S): Shiio, Isamu; Otsuka, Shinichiro; Ishii, Ryosuke;

Katsuya, Nobu; Iizuka, Hiroshi  
 CORPORATE SOURCE: Ajinomoto Co., Inc., Kawasaki, Japan  
 SOURCE: J. Gen. Appl. Microbiol. (Tokyo) (1963), 9, 23-30  
 DOCUMENT TYPE: Journal  
 LANGUAGE: Unavailable  
 TI **Microbial production of amino acids**  
 from hydrocarbons. I. Preliminary screening of glutamic acid-producing  
 bacteria  
 AB Various bacteria utilized kerosene, light oil, heavy oil, and liquid  
 paraffin as the only C source for growth and formation of L-glutamic acid  
 (I). The highest level of I (281 .gamma./ml.) was obtained from kerosene  
 by a strain of **Corynebacterium hydrocarboclastus**.

=> dis L15 1-13 ibib kwic

L15 ANSWER 1 OF 13 BIOSIS COPYRIGHT 2001 BIOSIS  
 ACCESSION NUMBER: 2001:314459 BIOSIS  
 DOCUMENT NUMBER: PREV200100314459  
 TITLE: Effect of gluconic acid as a secondary carbon source on  
 non-growing L-**lysine** producers cells of  
**Corynebacterium glutamicum**. Purification and  
 properties of 6-phosphogluconate dehydrogenase.  
 AUTHOR(S): Bianchi, Daniella; Bertrand, Olivier; Haupt, Karsten;  
 Coello, Nereida (1)  
 CORPORATE SOURCE: (1) Instituto de Biologia Experimental, Universidad  
 Central  
 deVenezuela, Caracas, 1041-A: ncoello@uole.com.ve  
 Venezuela  
 SOURCE: Enzyme and Microbial Technology, (June 7, 2001) Vol. 28,  
 No. 9-10, pp. 754-759. print.  
 ISSN: 0141-0229.  
 DOCUMENT TYPE: Article  
 LANGUAGE: English  
 SUMMARY LANGUAGE: English  
 TI Effect of gluconic acid as a secondary carbon source on non-growing L-  
**lysine** producers cells of **Corynebacterium glutamicum**.  
 Purification and properties of 6-phosphogluconate dehydrogenase.  
 AB We studied the production of L-**lysine** in **Corynebacterium**  
**glutamicum** ATCC 21543 non growing cells obtained by nutrient limitation.  
 Statistical analysis revealed significant differences in the L-  
**lysine** titers of glucose, gluconic acid or glucose-gluconic acid  
 cultures. Higher L-**lysine** titer obtained in batch cultures with  
 mixed carbon sources or gluconic acid alone were found to be associated  
 with a . . . dehydrogenase activity (6PGDH, E.C.1.1.1.44). This enzyme  
 is a pivotal enzyme within the hexose monophosphate pathway, and thus of  
 importance for L-**lysine** production. 6PGDH was purified and  
 characterized. The purified enzyme migrates as a single band on sodium  
 dodecyl sulfate-polyacrylamide gel electrophoresis. . .  
 IT . . .  
 Bioprocess Engineering; Methods and Techniques; Nutrition  
 IT Chemicals & Biochemicals  
 6-phosphogluconate dehydrogenase: amino acid sequence, analysis,  
 molecular properties, pH, purification; L-**lysine**:  
**microbial production**, yield; **amino**  
**acids**: analysis; carbon sources; gluconic acid: secondary  
 carbon source  
 ORGN . . .

Microorganisms; Irregular Nonsporing Gram-Positive Rods: Actinomycetes and Related Organisms, Eubacteria, Bacteria, Microorganisms; Microorganisms

ORGN Organism Name

Bacillus subtilis (Endospore-forming Gram-Positives);  
**Corynebacterium** glutamicum (Irregular Nonsporing Gram-Positive Rods): non-growing cells; Escherichia coli (Enterobacteriaceae); bacteria (Bacteria); microorganisms (Microorganisms)

ORGN Organism Superterms

Bacteria; Eubacteria; Microorganisms

RN 9001-82-5Q (6-PHOSPHOGLUCONATE DEHYDROGENASE)  
9073-95-4Q (6-PHOSPHOGLUCONATE DEHYDROGENASE)  
56-87-1 (L-**LYSINE**)  
526-95-4 (GLUCONIC ACID)

L15 ANSWER 2 OF 13 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER: 2001:174925 BIOSIS

DOCUMENT NUMBER: PREV200100174925

TITLE: MALDI-TOF MS for quantification of substrates and products in cultivations of **Corynebacterium** glutamicum.

AUTHOR(S): Wittmann, Christoph (1); Heinzle, Elmar

CORPORATE SOURCE: (1) Biochemical Engineering Institute, Saarland University,

66041, Saarbruecken; c.wittmann@rz.uni-sb.de Germany

SOURCE: Biotechnology and Bioengineering, (March 20, 2001) Vol. 72,

No. 6, pp. 642-647. print.

ISSN: 0006-3592.

DOCUMENT TYPE: Article

LANGUAGE: English

SUMMARY LANGUAGE: English

TI MALDI-TOF MS for quantification of substrates and products in cultivations

of **Corynebacterium** glutamicum.

AB The application of MALDI-TOF MS for the quantification of **lysine**, alanine, and glucose is described. The method is based on using stable isotopes as internal standards and allows fast, sensitive, . . . concentrations of the analytes between 10  $\mu$ M and 100 mM. The mean standard deviations from five replicates each were 4.3% (**lysine**), 3.7% (alanine), and 3.2% (glucose). In addition, sucrose could be measured by MALDI-TOF MS, but was not quantified due to. . .

IT Major Concepts

Biochemistry and Molecular Biophysics; Bioprocess Engineering; Methods and Techniques

IT Chemicals & Biochemicals

**amino acids: microbial production**

, quantitative analysis; products: quantitative analysis; substrates: quantitative analysis

L15 ANSWER 3 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 2000:761605 CAPLUS

DOCUMENT NUMBER: 134:99608

TITLE: Development and use of miniaturized parallel experiment technology for bioprocess development

AUTHOR(S): Altenbach-Rehm, Jutta

CORPORATE SOURCE: Institut fur Biotechnologie, Julich, JUL-3782, Germany

SOURCE: Ber. Forschungszent. Juelich (2000), Juel-3782, a-f,

DOCUMENT TYPE:

Report

LANGUAGE:

German

AB The fed-batch technique is nowadays the std. operation mode for high performance **microbial prodn.** processes. Shake flasks are widely used as simple bioreactors in batch process development. Because of uncontrolled changes in pH and reduced oxygen transfer rates parallel operated shake flasks can usually not be applied for microbial fed-batch process development. Due to the need to operate controlled stirred tank reactors the development of specific fermn. strategies is expensive and time consuming. To address these issues a miniature fed-batch technique was developed in cooperation with DASGIP mbH, Julich, and INFORS AG, Basel. Controlled batch and fed-batch fermns. can be performed in 16 parallel small scale bioreactors. The new technique allows the feeding of up to 4 different substrates and parallel pH

control

based on user-defined profiles. The feeding assembly is sterilized chem. using dimethylcarbonate. To overcome the limitations of shake flasks, small scale bubble columns were developed. Gas distribution is performed with a new type of sterile sparger. Transferring fed-batch fermns. from small scale bubble columns to a stirred tank reactor a scale up factor of 20 was achieved. *Escherichia coli* K12 was chosen to test the new

parallel

bioreactor technique. Compared to shake flask fermns. the cell concn.

was

50% higher due to efficient oxygen transfer. In fed-batch fermns. with pH-controlled substrate feeding up to 35 g/l DCW were achieved. For the 1st time, a parallel optimization of feeding profiles in parallel small scale fed-batch expts. with successful scale-up to a lab. bioreactor was performed. *Escherichia coli* BL 21 (DE3) pLysS produces the recombinant enzyme GDP-.alpha.-D-mannose-pyrophosphorylase after induction with isopropyl-.beta.-D-thiogalactoside (IPTG). Single induction with 0,5 mM IPTG resulted in a low specific enzyme activity of 1,6 U/g DCW (dry cell wt.). For the optimization of enzyme expression a genetic algorithm was used. A final enzyme activity of 111 U/g DCW was achieved for optimal substrate and inducer feeding profiles. To demonstrate the advantages of this new parallel bioreactor technique different strains of industrial relevance were investigated. **Corynebacterium glutamicum**, *Staphylococcus carnosus* and *Ashbya gossypii*.

ST bioreactor bubble column fed batch fermn; GDP mannose pyrophosphorylase bubble column fed batch; *Staphylococcus* bubble column fed batch fermn; isoleucine bubble column fed batch **Corynebacterium**; riboflavin bubble column fed batch *Ashbya*

IT **Amino acids**, biological studies

RL: BPR (Biological process); BIOL (Biological study); PROC (Process) (amino acid consumption in riboflavin prodn. by *Ashbya gossypii* in parallel bubble columns with fed-batch technique)

IT **Corynebacterium glutamicum**

(L-isoleucine prodn. by **Corynebacterium glutamicum** in parallel bubble columns with fed-batch technique)

IT 56-40-6, Glycine, biological studies 56-41-7, Alanine, biological studies 56-45-1, Serine, biological studies 56-84-8, Aspartic acid, biological studies 56-85-9, Glutamine, biological studies 56-86-0, Glutamic acid, biological studies 56-87-1, **Lysine**, biological studies 60-18-4, Tyrosine, biological studies 63-68-3, Methionine, biological studies 63-91-2, Phenylalanine, biological studies

72-18-4,

Valine, biological studies 72-19-5, Threonine, biological studies  
 73-22-3, Tryptophane, biological studies  
 RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
 (amino acid consumption in riboflavin prodn. by *Ashybya gossypii* in  
 parallel bubble columns with fed-batch technique)  
 IT 73-32-5P, L-Isoleucine, biological studies  
 RL: BMF (Bioindustrial manufacture); BPR (Biological process); BIOL  
 (Biological study); PREP (Preparation); PROC (Process)  
 (L-isoleucine prodn. by *Corynebacterium glutamicum* in  
 parallel bubble columns with fed-batch technique, amino acid  
 consumption in riboflavin prodn. by *Ashybya gossypii*)  
 IT 61-90-5, L-Leucine, biological studies  
 RL: BPR (Biological process); MFM (Metabolic formation); BIOL (Biological  
 study); FORM (Formation, nonpreparative); PROC (Process)  
 (L-isoleucine prodn. by *Corynebacterium glutamicum* in  
 parallel bubble columns with fed-batch technique, amino acid  
 consumption in riboflavin prodn. by *Ashybya gossypii*)

L15 ANSWER 4 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1999:244776 CAPLUS

DOCUMENT NUMBER: 130:266420

TITLE: Method for **microbial production** of  
**amino acids** of the aspartate and/or  
 glutamate family and agents which can be used in said  
 method

INVENTOR(S): Eikmanns, Bernd; Peters-Wendisch, Petra; Sahm,  
 Hermann

PATENT ASSIGNEE(S): Forschungszentrum Julich G.m.b.H., Germany

SOURCE: PCT Int. Appl., 40 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9918228	A2	19990415	WO 1998-EP6210	19980930
WO 9918228	A3	19990520		
W:	AU, BR, CA, CN, HU, ID, JP, KR, MX, RU, SK, US			
RW:	AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE			
DE 19831609	A1	19990415	DE 1998-19831609	19980714
AU 9911482	A1	19990427	AU 1999-11482	19980930
EP 1015621	A2	20000705	EP 1998-954301	19980930
R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI			
BR 9813021	A	20000815	BR 1998-13021	19980930
PRIORITY APPLN. INFO.:			DE 1997-19743894 A	19971004
			DE 1998-19831609 A	19980714
			WO 1998-EP6210 W	19980930

TI Method for **microbial production** of **amino  
 acids** of the aspartate and/or glutamate family and agents which  
 can be used in said method

AB The invention relates to a method for **microbial prodn.**  
 of **amino acids** of the aspartate and/or glutamate  
 family in which the pyruvate carboxylase activity is increased by  
 genetically changing the enzyme and/or the pyruvate carboxylase gene

expression of a microorganism which produces the corresponding amino acid.

In addn., the invention relates to a pyruvate carboxylase gene and addnl. agents which can be used in the inventive method.

ST amino acid fermn **Corynebacterium** pyruvate carboxylase genetic engineering

IT **Corynebacterium** glutamicum  
Fermentation  
(microbial prodn. of amino acids  
of the aspartate and/or glutamate family and agents which can be used in said method)

IT **Amino acids**, preparation  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(microbial prodn. of amino acids  
of the aspartate and/or glutamate family and agents which can be used in said method)

IT Genetic engineering  
(microbial prodn. of amino acids  
of the aspartate and/or glutamate family and modification of **Corynebacterium** pyc gene in said method)

IT Genes (microbial)  
RL: BPR (Biological process); BIOL (Biological study); PROC (Process)  
(pyc; microbial prodn. of amino acids of the aspartate and/or glutamate family and modification of **Corynebacterium** pyc gene in said method)

IT 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, preparation 72-19-5P, L-Threonine, preparation 74-79-3P, L-Arginine, preparation 672-15-1P, L-Homoserine  
RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(microbial prodn. of amino acids  
of the aspartate and/or glutamate family and agents which can be used in said method)

IT 9014-19-1, Pyruvate carboxylase 204116-18-7 208541-81-5  
RL: BSU (Biological study, unclassified); BIOL (Biological study)  
(microbial prodn. of amino acids  
of the aspartate and/or glutamate family and agents which can be used in said method)

L15 ANSWER 5 OF 13 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER: 1999:175414 BIOSIS

DOCUMENT NUMBER: PREV199900175414

TITLE: Cloning of the transketolase gene and the effect of its dosage on aromatic amino acid production in **Corynebacterium** glutamicum.

AUTHOR(S): Ikeda, M. (1); Okamoto, K.; Katsumata, R.

CORPORATE SOURCE: (1) Technical Research Laboratories, Kyowa Hakko Kogyo Co.,

SOURCE: Ltd., Hofu, Yamaguchi, 747-8522 Japan  
Applied Microbiology and Biotechnology, (Feb., 1999) Vol. 51, No. 2, pp. 201-206.  
ISSN: 0175-7598.

DOCUMENT TYPE: Article

LANGUAGE: English

TI Cloning of the transketolase gene and the effect of its dosage on aromatic

amino acid production in **Corynebacterium** glutamicum.

AB. . . enzyme of the non-oxidative pentose phosphate pathway. The effect of

its overexpression on aromatic amino acid production was investigated in **Corynebacterium glutamicum**, a typical amino-acid-producing organism. For this purpose, the transketolase gene of the organism was cloned on the basis of. . . as a protein of approximately 83kDa in proportion to the copy numbers. Introduction of the plasmids into a tryptophan and **lysine** co-producer resulted in copy-dependent increases in tryptophan production along with concomitant decreases in **lysine** production. Furthermore, the presence of the gene in high copy numbers enabled tyrosine, phenylalanine and tryptophan producers to accumulate 5%-20% more aromatic **amino acids**. These results indicate that overexpressed transketolase activity operates to redirect the glycolytic intermediates toward the nonoxidative pentose phosphate pathway in. . .

IT Major Concepts

Bioprocess Engineering; Enzymology (Biochemistry and Molecular Biophysics); Molecular Genetics (Biochemistry and Molecular

Biophysics)

IT Chemicals & Biochemicals

aromatic **amino acids: microbial**

**production**; transketolase [EC 2.2.1.1]; **Corynebacterium**

transketolase gene (Irregular Nonsporing Gram-Positive Rods)

L15 ANSWER 6 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1995:674124 CAPLUS

DOCUMENT NUMBER: 123:54314

TITLE: Enhancement of reduced NADP production for enhanced **microbial production** of biochemicals

INVENTOR(S): Kojima, Hiroyuki; Totsuka, Kazuhiko

PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan

SOURCE: PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9511985	A1	19950504	WO 1994-JP1791	19941026
W: AU, BR, CA, CN, CZ, HU, JP, KR, PL, RU, SK, US, VN				
RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
CA 2175042	AA	19950504	CA 1994-2175042	19941026
AU 9480026	A1	19950522	AU 1994-80026	19941026
AU 687458	B2	19980226		
EP 733712	A1	19960925	EP 1994-931158	19941026
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT,				
SE				
BR 9407907	A	19961126	BR 1994-7907	19941026
HU 74840	A2	19970228	HU 1996-1085	19941026
ZA 9503350	A	19961025	ZA 1995-3350	19950425
US 5830716	A	19981103	US 1996-619521	19960429
CN 1139956	A	19970108	CN 1994-194707	19961026
PRIORITY APPLN. INFO.:			JP 1993-270828	19931028
			WO 1994-JP1791	19941026

TI Enhancement of reduced NADP production for enhanced **microbial production** of biochemicals



AB The productivity of such substances as L-**amino acids**, antibiotics, vitamins, growth factors and physiol. active substances in the fermn. using a microorganism is improved by improving the productivity of reduced NADP in the cells of the microorganisms. Construction of pMW::THY contg. the Escherichia coli transhydrogenase gene, and introduction of the plasmid into the L-threonine-producing Escherichia coli B-3996 were shown. The recombinant E. coli B-3996 produced L-threonine .apprx.10% higher than did the parental strain.

IT **Corynebacterium** glutamicum  
Escherichia coli  
Fermentation  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT **Amino acids**, preparation  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT Plasmid and Episome  
(pHSG::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT Plasmid and Episome  
(pMW::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT Plasmid and Episome  
(pSU::THY; enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT 9014-18-0, Nicotinamide nucleotide transhydrogenase  
RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-**Lysine**, preparation 61-90-5P, L-Leucine, preparation 63-91-2P, L-Phenylalanine, preparation 72-18-4P, L-Valine, preparation 72-19-5P, L-Threonine, preparation 73-32-5P, L-Isoleucine, preparation  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

IT 53-59-8P, NADP  
RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)  
(reduced; enhancement of reduced NADP prodn. for enhanced **microbial prodn.** of biochems.)

L15 ANSWER 7 OF 13 SCISEARCH COPYRIGHT 2001 ISI (R)  
ACCESSION NUMBER: 95:184306 SCISEARCH  
THE GENUINE ARTICLE: QK574  
TITLE: METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM  
**CORYNEBACTERIUM-GLUTAMICUM**  
AUTHOR: SAHM H (Reprint); EGDELING L; EIKMANN B; KRAMER R  
CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL,  
D-52425 JULICH, GERMANY (Reprint)  
COUNTRY OF AUTHOR: GERMANY  
SOURCE: FEMS MICROBIOLOGY REVIEWS, (FEB 1995) Vol. 16, No. 2-3,

pp. 243-252.  
ISSN: 0168-6445.

DOCUMENT TYPE: Article; Journal  
FILE SEGMENT: LIFE  
LANGUAGE: ENGLISH  
REFERENCE COUNT: 36

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

TI METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM **CORYNEBACTERIUM**  
-GLUTAMICUM

AB The Gram-positive bacterium **Corynebacterium** glutamicum is used for the industrial production of **amino acids**, e.g. of L-glutamate and L-**lysine**. In the last 10 years, genetic engineering and amplification of relevant structural genes have become fascinating methods for the construction of strains with desired genotypes. By cloning and expressing the various genes of the L-**lysine** pathway in *C. glutamicum* we could demonstrate that an increase of the flux of L-aspartate semialdehyde to L-**lysine** could be obtained in strains with increased dehydrodipicolinate synthase activity. By combined overexpression of deregulated aspartate kinase and dihydrodipicolinate synthase, the L-**lysine** secretion could be increased (10-20%). Recently we detected that in *C. glutamicum* two pathways exist for the synthesis of DL-diaminopimelate and L-**lysine**. Mutants defective in one pathway are still able to synthesize enough L-**lysine** for growth, but the L-**lysine** secretion is reduced to 50-70%. Using NMR spectroscopy, we could calculate

how much of the L-**lysine** secreted into the medium is synthesized via each pathway. Amplification of the feedback inhibition-insensitive homoserine dehydrogenase and homoserine kinase in a high L-**lysine** overproducing strain enabled channelling of the carbon flow from the intermediate aspartate semialdehyde towards homoserine, resulting in a high accumulation. . . . acid overproduction, the secretion into the culture medium also has to be noted. Recently it could be demonstrated that L-glutamate, L-**lysine** and L-isoleucine are not secreted via passive diffusion but via specific active carrier systems. Analysis of **lysine**-overproducing *C. glutamicum* strains indicates that this secretion carrier has a strong influence on the overproduction of this amino acid. Thus, . . .

ST Author Keywords: **CORYNEBACTERIUM** GLUTAMICUM; AMINO ACID  
PRODUCTION; METABOLIC DESIGN; L-**LYSINE**; L-THREONINE;  
L-ISOLEUCINE

STP KeyWords Plus (R): L-THREONINE; L-**LYSINE**; BREVIBACTERIUM-  
LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; **MICROBIAL**  
**PRODUCTION**; RESISTANT MUTANTS; SPLIT PATHWAY; BIOSYNTHESIS;  
ISOLEUCINE; GENES

L15 ANSWER 8 OF 13 SCISEARCH COPYRIGHT 2001 ISI (R)

ACCESSION NUMBER: 96:7187 SCISEARCH

THE GENUINE ARTICLE: TJ545

TITLE: METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM  
**CORYNEBACTERIUM**-GLUTAMICUM

AUTHOR: SAHM H (Reprint)

CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL,  
D-52425 JULICH, GERMANY (Reprint)

COUNTRY OF AUTHOR: GERMANY

SOURCE: FOLIA MICROBIOLOGICA, (1995) Vol. 40, No. 1, pp. 23-30.  
ISSN: 0015-5632.

DOCUMENT TYPE: Article; Journal

FILE SEGMENT: LIFE; AGRI  
LANGUAGE: ENGLISH  
REFERENCE COUNT: 26

\*ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS\*

TI METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM  
**CORYNEBACTERIUM-GLUTAMICUM**

AB The Gram-positive bacterium **Corynebacterium glutamicum** is used for the industrial production of **amino acids**, e.g. of L-glutamate and L-**lysine**. By cloning and expressing the various genes of the L-**lysine** pathway in **C. glutamicum** we could demonstrate that an increase of the flux of L-4-aspartaldehyde to L-**lysine** could be obtained in strains with increased dihydro-dipicolinate synthase activity. Recently we detected that in **C. glutamicum** two pathways exist for the synthesis of DL-2,6-diaminopimelate and L-**lysine**. Mutants defective in one pathway are still able to synthesize enough L-**lysine** for growth but the L-**lysine** secretion is reduced to 50-70 %. Using NMR-spectroscopy we could calculate

how much of the L-**lysine** secreted into the medium is synthesized via the one and the other pathway. Amplification of the feedback-inhibition-insensitive-homoserine dehydrogenase and homoserine kinase in a high L-**lysine**-overproducing strain made it possible to channel the carbon now from the intermediate 4-aspartaldehyde toward homoserine, resulting in a high. . .

STP KeyWords Plus (R): L-THREONINE; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; **MICROBIAL PRODUCTION; LYSINE**  
BIOSYNTHESIS; RESISTANT MUTANTS; SPLIT PATHWAY; GENES; AMPLIFICATION; FERMENTATION

L15 ANSWER 9 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1985:436156 CAPLUS

DOCUMENT NUMBER: 103:36156

TITLE: Optimization of amino acid production by automatic self-tuning digital control of redox potential

AUTHOR(S): Radjai, Mohammad K.; Hatch, Randolph T.; Cadman, Theodore W.

CORPORATE SOURCE: Dep. Chem. Nucl. Eng., Univ. Maryland, College Park, MD, 20742, USA

SOURCE: Biotechnol. Bioeng. Symp. (1984), 14(Symp. Biotechnol.

Fuels Chem., 6th), 657-79

CODEN: BIBSBR; ISSN: 0572-6565

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The **microbial prodn.** of homoserine [672-15-1], **lysine** [56-87-1], and valine [72-18-4] by an auxotrophic mutant of **Corynebacterium glutamicum** was investigated in a 16-L batch fermentor. Closed-loop digital control of redox potential was

implemented

using proportional-integral (PI) control of agitation rates. Due to the nonlinearity of the system, the PI controller parameters had to be changed

during the course of the ferms. An automatic, self-tuning algorithm was developed for stable control of redox potential. This permitted exptl. optimization of total and selective amino acid prodn. Total amino acid yields of 35% from glucose were achieved compared to 23% reported in the literature for the same fermn.

ST amino acid fermn redox potential control; optimization simulation

homoserine **lysine** valine fermn  
 IT **Corynebacterium** glutamicum  
 (amino acid manuf. with, optimization and redox potential control in)  
 IT **Amino acids**, preparation  
 RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP  
 (Preparation)  
 (manuf. of, by fermn.)

L15 ANSWER 10 OF 13 MEDLINE  
 ACCESSION NUMBER: 79079819 MEDLINE  
 DOCUMENT NUMBER: 79079819 PubMed ID: 727028  
 TITLE: **Microbial production** of essential amino  
 acid with **Corynebacterium** glutamicum mutants.  
 AUTHOR: Nakayama K; Araki K; Kase H  
 SOURCE: ADVANCES IN EXPERIMENTAL MEDICINE AND BIOLOGY, (1978) 105  
 649-61.  
 Journal code: 2LU; 0121103. ISSN: 0065-2598.  
 PUB. COUNTRY: United States  
 Journal; Article; (JOURNAL ARTICLE)  
 LANGUAGE: English  
 FILE SEGMENT: Priority Journals  
 ENTRY MONTH: 197902  
 ENTRY DATE: Entered STN: 19900314  
 Last Updated on STN: 19970203  
 Entered Medline: 19790212

TI **Microbial production** of essential amino acid with  
**Corynebacterium** glutamicum mutants.  
 AB **Amino acids** produced by microbial process are  
 generally L-forms. The stereospecificity of the **amino**  
**acids** produced by fermentation makes the process advantageous  
 compared with synthetic process. Microorganisms employed in microbial  
 process for amino acid production are divided into 4 classes; wild-type  
 strain, auxotrophic mutant, regulatory mutant and auxotrophic regulatory  
 mutant. Using such mutants of **Corynebacterium** glutamicum, all  
 the essential **amino acids** but L-methionine are now  
 being produced by "direct fermentation" from cheap carbon sources such as  
 carbohydrate materials or acetic acid.

CT **\*Amino Acids, Essential: BI, biosynthesis**  
**\*Corynebacterium: ME, metabolism**

Fermentation  
 Kinetics  
 Leucine: BI, biosynthesis  
**Lysine: BI, biosynthesis**  
 Mutation  
 Phenylalanine: BI, biosynthesis  
 Species Specificity  
 Stereoisomerism  
 Threonine: BI, biosynthesis  
 Tryptophan: BI, biosynthesis

RN 3617-44-5 (Phenylalanine); **56-87-1 (Lysine)**; 7005-03-0  
 (Leucine); 72-19-5 (Threonine); 73-22-3 (Tryptophan)  
 CN 0 (**Amino Acids, Essential**)

L15 ANSWER 11 OF 13 CAPLUS COPYRIGHT 2001 ACS  
 ACCESSION NUMBER: 1970:508239 CAPLUS  
 DOCUMENT NUMBER: 73:108239  
 TITLE: **Microbial production** of  
 L-threonine

INVENTOR(S): Nakayama, Kiyoshi; Kase, Hiroshi  
 PATENT ASSIGNEE(S): Kyowa Fermentation Industry Co. Ltd.  
 SOURCE: Ger. Offen., 22 pp.  
 CODEN: GWXXBX  
 DOCUMENT TYPE: Patent  
 LANGUAGE: German  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	DE 1817666	A	19700827	DE 1968-1817666	19681224

TI **Microbial production of L-threonine**  
 AB Various microorganisms, e.g. Aerobacter [Enterobacter] aerogenes, Serratia marcescens, or Arthrobacter paraffineus, cultured for producing L-threonine required 2 or 3 of the **amino acids** isoleucine, methionine, **lysine**, or diaminopimelic acid. The microorganisms were cultured aerobically in an aq. medium contg. the optimal (or less) amts. of the required **amino acids**. Thus, E. aerogenes NM-IS-5 (ATCC 21,215) was cultured 96 hr at 30.degree. in medium contg. glucose 5, (NH4)2SO4 1.4, KH2PO4 0.05, K2HPO4 0.05, MgSO4.7H2O 0.025, FeSO4.7H2O 0.001, MnSO4.4H2O 0.001, and CaCO3 2% and isoleucine 50, methionine 100, and diaminopimelic acid 200 mg/l. to give 7.8 g L-threonine/l.  
 ST **microbial prodn** threonine; threonine **microbial prodn**; Aerobacter threonine fermn; amino acid prodn fermn  
 IT **Corynebacterium** (glutamicum, threonine manuf. by)

L15 ANSWER 12 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1967:514494 CAPLUS

DOCUMENT NUMBER: 67:114494

TITLE: **Microbial production of amino acids** from hydrocarbons. III. L-Ornithine production by an arginine auxotrophic mutant of **Corynebacterium** hydrocarboclastus  
 AUTHOR(S): Ishu, Ryosuke; Ishii, Ryosuke; Shio, Isamu  
 CORPORATE SOURCE: Ajinomoto Co., Inc., Kawasaki, Japan  
 SOURCE: J. Gen. Appl. Microbiol. (1967), 13(3), 3303-12  
 CODEN: JGAMA9

DOCUMENT TYPE: Journal

LANGUAGE: English

TI **Microbial production of amino acids** from hydrocarbons. III. L-Ornithine production by an arginine auxotrophic mutant of **Corynebacterium** hydrocarboclastus  
 AB cf. CA 67: 89718u. The arginine auxotrophic mutant strain RN-362 of C. hydrocarboclastus R-7 was used to study L-ornithine production from hydrocarbons, in a fermentation medium contg. various n-alkanes. L-Ornithine production required L-arginine at the optimum level of 0.5-1.0 g./l. of medium; an excess inhibited the biosynthesis of L-ornithine. (NH4)2HPO4 was the best source of N and, at 2% in a neutral to slightly acidic pH, gave the highest level of L-ornithine production and cell growth; NH4OAc, KNO3, and (NH4)2CO3 proved less suitable because of a drop in pH along with the accumulation of a large amt. of .alpha.-ketoglutaric acid, pyruvic acid, and proline in the growth medium. Of 17 C sources,

n-tetradecane best supported cell growth and L-ornithine production and the other C13-C17 n-alkanes did so moderately, while kerosene and light oil produced good cell growth but only a small amt. of L-ornithine.

Addn.

of 3 g. yeast ext. and 0.5 g. L-arginine-HCl to 1 l. of medium enhanced L-ornithine production. A similar effect was achieved by replacing the yeast ext. with various **amino acids** at 0.01% in the medium. L-Methionine was most effective for the production of L-ornithine, while L-lysine, L-cysteine, L-cystine, L-histidine, and L-phenylalanine were less so, in decreasing order. **Amino acids** enhance L-ornithine production by stimulating hydrocarbon oxidn. and cell growth.

ST HYDROCARBONS USE BACTERIA; BACTERIA HYDROCARBONS USE; ALKANES USE BACTERIA; **AMINO ACIDS** PRODN HYDROCARBONS; ORNITHINE PRODN HYDROCARBONS; PARAFFINS UTILIZATION BACTERIA

IT **Corynebacterium**

(hydrocarboclastus, ornithine formation from hydrocarbons by)

IT Hydrocarbons, biological studies

RL: BIOL (Biological study)

(ornithine formation from, by **Corynebacterium** hydrocarboclastus)

IT 70-26-8

RL: FORM (Formation, nonpreparative)

(formation of, from hydrocarbons by **Corynebacterium** hydrocarboclastus)

L15 ANSWER 13 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1967:489718 CAPLUS

DOCUMENT NUMBER: 67:89718

TITLE:

**Microbial production of**

**amino acids** from hydrocarbons. II.

Isolationf good hycarbon utilizers and amino acid production by their auxotrophs

AUTHOR(S): Ishii, Ryosuke; Otsuka, Shinichiro; Shiio, Isamu

CORPORATE SOURCE: Central Res. Labs., Ajinomoto Co., Inc., Kawasaki, Japan

SOURCE: J. Gen. Appl. Microbiol. (1967), 13(2), 217-25

CODEN: JGAMA9

DOCUMENT TYPE: Journal

LANGUAGE: English

TI **Microbial production of amino acids**

from hydrocarbons. II. Isolationf good hycarbon utilizers and amino acid

production by their auxotrophs

AB cf. CA 59: 14313h. Nine microorganisms, which showed good growth on long-chain aliphatic hydrocarbons, were isolated by an enrichment culture method, followed by a single colony isolation technique. They included 5 strains of *Alcaligenes marshallii*, 2 strains of **Corynebacterium** hydrocarboclastus, and 2 strains of yeast. Various auxotrophic mutants were derived from these microorganisms. The mutants accumulated the following **amino acids** from aliphatic hydrocarbons; L-ornithine, L-valine, L-glutamic acid, L-leucine, L-tyrosine, L-alanine, L-proline, L-aspartic acid, and L-lysine.

ST BACTERIA AMINO ACID PRODN; AMINO ACID PRODN BACTERIA; HYDROCARBONS **AMINO ACIDS**; ALIPHATICS BACTERIA METAB

IT **Corynebacterium**

(hydrocarboclastus, amino acid fermentation of hydrocarbons by)

IT **Amino acids**, preparation

RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP  
(Preparation)  
(manuf. of, by fermentation of hydrocarbons)

=> dis his

(FILE 'HOME' ENTERED AT 18:56:31 ON 19 OCT 2001)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, SCISEARCH' ENTERED AT 18:57:06 ON  
19 OCT 2001

L1 2893 S MICROBIAL (W) PRODUCTION  
L2 190 S L1 AND (AMINO (W) ACIDS)  
L3 28 S L2 AND CORYNEBACTERIUM  
L4 13 L3 AND LYSINE  
L5 0 L3 AND ((EXPORT) (W) (GENE OR CARRIER))  
L6 0 S L3 AND EXPORT (W) GENE  
L7 166 S EXPORT (W) GENE  
L8 0 S L3 AND L7  
L9 0 S L3 (P) L7  
L10 0 S L2 AND L7  
L11 0 S L2 AND EXPORT (W) GENE  
L12 61 S L7 AND MICROB?  
L13 1 S L12 AND CORYNEBACTERIUM  
L14 26 DUP REM L3 (2 DUPLICATES REMOVED)  
L15 13 DUP REM L4 (0 DUPLICATES REMOVED)

=> log off y

\* \* \* \* \*

Dear valued customer,

Your feedback is important to us. Would you kindly take a moment to  
complete our survey? This survey will only take about 5-10 minutes to  
complete. Your responses will be kept confidential and will help us  
improve STN Express with Discover! for your future use. Please click  
on the following link to access the survey.

<http://www.cas.org/ONLINE/STN/ExpressSurveyForm.html?LOGINID=SSSPTA1600RXM>

\* \* \* \* \*

STN INTERNATIONAL LOGOFF AT 19:32:29 ON 19 OCT 2001